

Application No.: 08/466,554

Amendment dated February 16, 2005

Reply to Office Action mailed August 16, 2004

## Amendments to the Claims:

This listing of claims will replace all prior versions, and listings of claims in the application:

## Listing of Claims:

1-41. (Canceled)

42. (Previously Presented) A method for screening a compound to determine its ability to alter the amount of an Aβ (x-≥ 41) peptide in a cerebral spinal fluid sample comprising:

measuring a first amount of one or more soluble A $\beta$  (x- $\geq$  41) peptides in the cerebral spinal fluid sample of a non-human animal model that expresses amyloid- $\beta$  precursor protein (APP) in the brain and processes it to the one or more soluble A $\beta$  peptides;

administering the compound to the non-human animal model;

measuring a second amount of the one or more soluble  $A\beta$  peptides in the cerebral spinal fluid sample of the non-human animal model; and

comparing the first amount with the second amount,

the difference indicating whether the compound increases, decreases, or leaves unchanged the amount of soluble  $A\beta$  ( $x \ge 41$ ) in the cerebral spinal fluid sample, wherein the amount of  $A\beta(x \ge 41)$  is measured by exposing the cerebral spinal fluid to an antibody or fragment thereof specific for an epitope of  $A\beta(x \ge 41)$  but that does not cross react with an epitope of  $A\beta(x \le 40)$ .

43. (Previously Presented) The method of claim 42, wherein the non-human animal model is a rodent model.



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- 44. (Previously Presented) The method of claim 43, wherein the rodent model is a mouse model.
- 45. (Currently Amended) The method of claim 42, wherein the non-human animal model is a transgenic rodentanimal model having an expression cassette that drives the expression of a sequence which encodes the Swedish mutation of an amyloid precursor protein (APP)APP gene.
  - 46. (Cancel)
  - 47. (Canceled)
- 48. (Previously Presented) The method of claim 42, wherein the antibody of fragment thereof binds to  $A\beta(x \ge 41)$  is  $A\beta(x-42)$ .

49-50. (Canceled)

- 51. (Previously Presented) The method of claim 42, wherein antibody or fragment thereof is an antibody recognizing an epitope on Aβ having amino acid residues 33-42.
- 52. (Previously Presented) The method of claim 51, wherein the antibody or fragment thereof is an antibody that will specifically bind to A $\beta$  (x- $\geq$  41).
  - 53. (Canceled)
- 54. (Previously Presented) The method of claim 51, wherein the antibody or fragment thereof is an antibody that will specifically bind to A $\beta$  (x- $\geq$  43).